

BREAST CANCER

RARRES3—suppressing metastases to the lung in breast cancer

About 90% of deaths due to breast cancer are caused by distant metastases. Oestrogen receptor (ER)-negative breast cancer tumours account for 20–30% of breast cancer cases, which primarily metastasize to the lungs. A group led by Roger Gomis has recently demonstrated that loss of function of *RARRES3*, a retinoic acid responsive gene, in ER-negative breast cancer cells stimulates their invasive capacity and promotes metastasis to the lung.

Gomis explains, “the transformation of a tumour cell into a metastatic cell is not just about acquiring capacities, but equally important is the loss of certain genes, such as *RARRES3*.” The researchers combined studies in mouse models and cell lines, and ultimately validated their findings in 580 samples from primary breast tumours, demonstrating that *RARRES3* is suppressed in ER-negative breast

tumours. Furthermore, *RARRES3* loss is associated with a lack of differentiation capacity, which facilitates the initiation of metastases. Therefore, *RARRES3* loss could provide a marker to identify patients with greater susceptibility to lung metastasis.

“We are currently exploring the potential clinical implications of our findings,” says Gomis. However, the researchers believe that some patients may benefit from treatment with retinoic acid after the removal of the original tumour, to induce *RARRES3* expression, enhance cell differentiation and prevent metastases.

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